

Figure 1

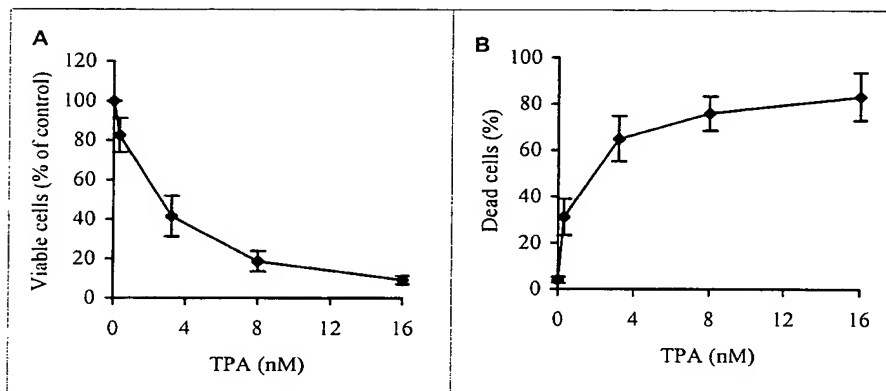


Figure 2

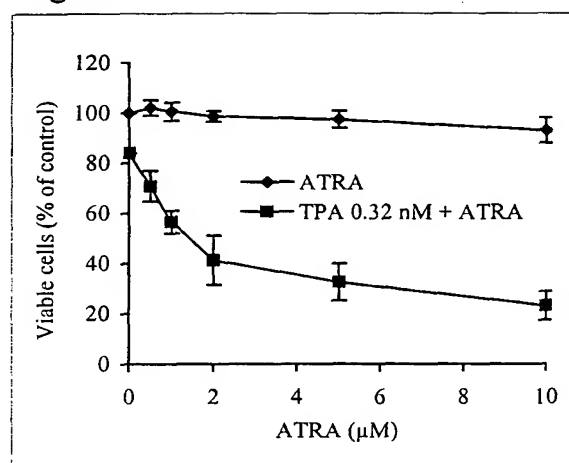


Figure 3

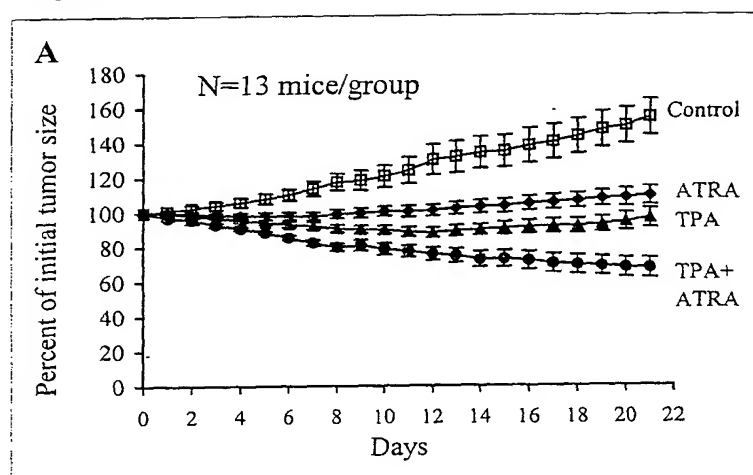


Figure 4

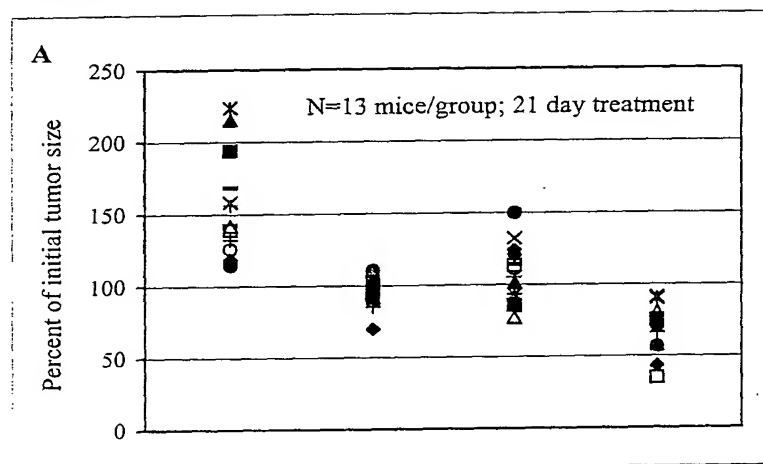


Figure 5

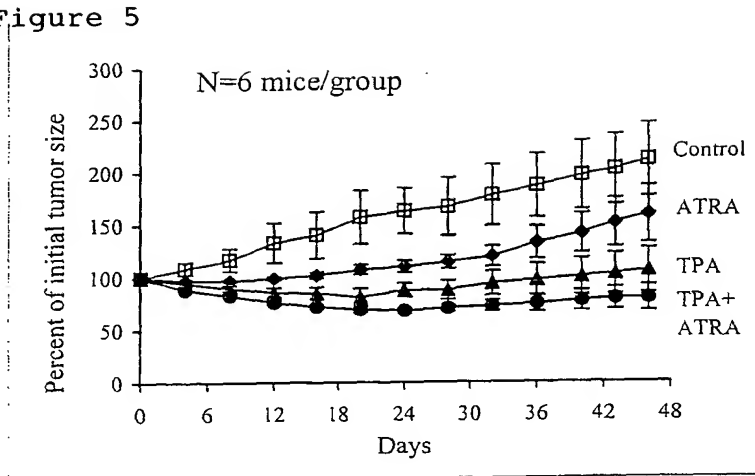
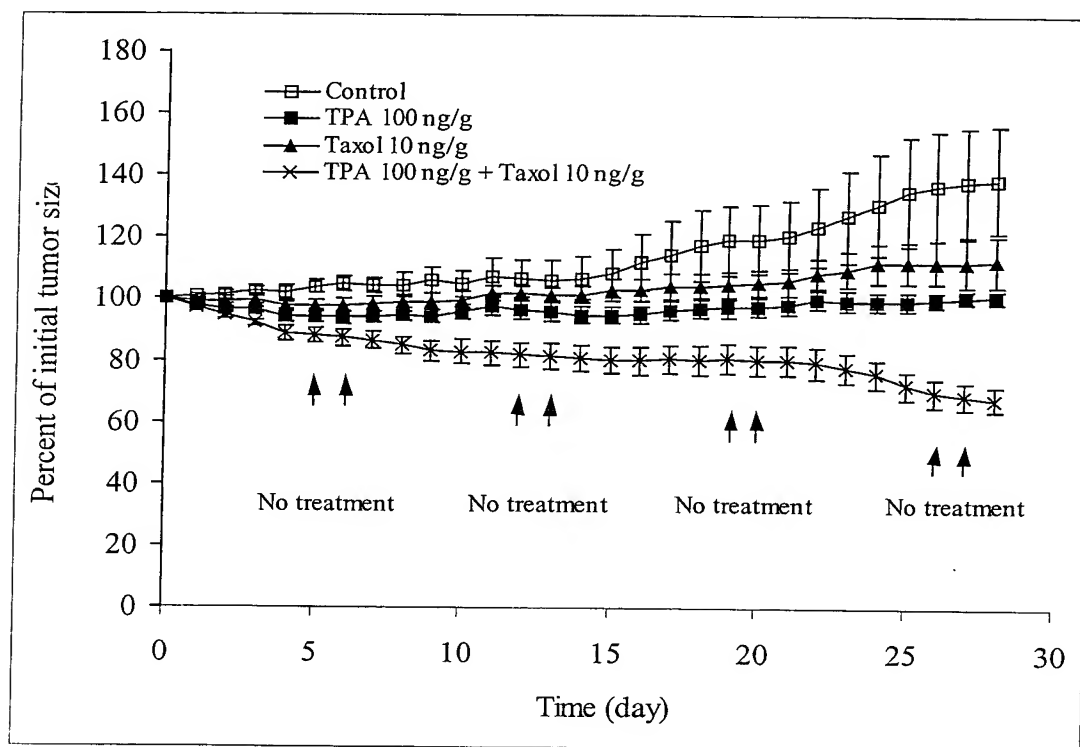


FIGURE 6

Effect of i.p. injections of TPA or Taxol alone or in combination on the growth of LNCaP tumors in NCr immunodeficient mice.



Male NCr immunodeficient mice were injected subcutaneously with LNCaP cells in matragel. After 4-6 weeks, mice with tumors (0.65-1 cm long and 0.65-1 cm wide) were randomly assigned to 4 groups (6 mice per group). Animals in group 1 received i.p. injections of vehicle (5 μ l/g body weight), animals in group 2 received i.p. injections of TPA (100 ng/g; 5 μ l vehicle/g), animals in group 3 received i.p. injections of Taxol (10 ng/g; 5 μ l vehicle/g) and animals in group 4 received i.p. injections of TPA (100 ng/g) in combination with Taxol (10 ng/g) in 5 μ l vehicle/g once a day for 5 days followed by a 2-day intermission. The mice received treatment for 28 days. The vehicle (5 μ l/g body weight) consists of propylene glycol, polysorbate 80, benzyl alcohol, ethanol and water (40: 0.5: 1: 10: 48.5). Tumor size (length x width) was measured and expressed as percent of initial size. Each value represents the mean \pm S.E.

FIGURE 7

Effects of TPA and Taxol alone or in combination on growth, apoptosis and cells cycle distribution in LNCaP cells

Treatment	No. of viable cells (1x10 ⁵ /ml)	Percent of cells in			% Apoptotic cells
		G0/G1	S	G2/M	
<u>Treatment for 48 hr</u>					
Control	8.9	76.5	16.1	5.2	1.6
Ethanol	10.0	76.8	16.4	4.6	1.8
TPA 1 ng/ml	3.7	77.8	13.0	5.5	10.1
Taxol 5ng/ml	4.6	73.0	12.8	6.4	12.9
TPA 1ng/ml + Taxol 5ng/ml	1.3	58.6	12.6	5.3	28.3
<u>Treatment for 96 hr</u>					
Control	10.6	85.7	7.5	5.2	1.2
Ethanol	12.3	83.6	7.9	6.2	1.9
TPA 1 ng/ml	3.2	66.7	12.3	9.5	11.2
Taxol 5ng/ml	2.5	49.7	12.4	8.7	28.4
TPA 1ng/ml + Taxol 5ng/ml	0.4	29.9	10.1	8.0	50.8

LNCaP cells were seeded at a density of $1.0 \times 10^5/\text{ml}$ in 60 mm culture dishes and incubated for 24 hr. The cells were then treated once with TPA (1 ng/ml) and Taxol (5 ng/ml) alone or in combination for 48 or 96 hr. Cell proliferation, apoptosis and cell cycle distribution were examined.